

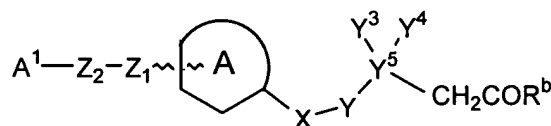
II. AMENDMENTS TO THE CLAIMS

Please amend the claims as follows:

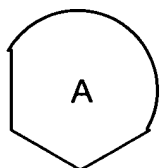
Claims 1-65 (Previously Cancelled)

Claims 66-71 (Cancelled)

Claim 71 (Original) A compound of the Formula I



or a pharmaceutically acceptable salt thereof, wherein

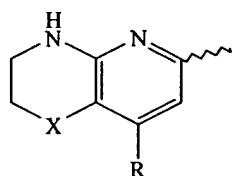


is a thiazole or isoxazole, optionally substituted with one or more substituent selected from the group consisting of alkyl, haloalkyl, aryl, heteroaryl, halogen, alkoxyalkyl, aminoalkyl, hydroxy, nitro, alkoxy, hydroxyalkyl, thioalkyl, amino, alkylamino, arylamino, alkylsulfonamide, acyl, acylamino, alkylsulfone, sulfonamide, allyl, alkenyl, methylenedioxy, ethylenedioxy, alkynyl, carboxamide, cyano, and $-(CH_2)_mCOR$;

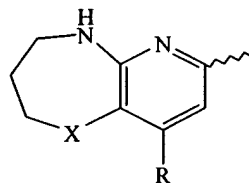
m is 0-2;

R is selected from the group consisting of hydroxy, alkoxy, alkyl, amino and sulfone;

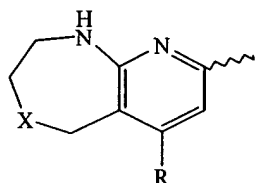
A¹ is selected from the group consisting of



X = CH₂, O, S, SO₂, CO,
CF₂, CMe₂
R = H, Me, OMe, OH, NR₂



X = CH₂, O, S, SO₂, CO,
CF₂, CMe₂
R = H, Me, OMe, OH, NR₂ and



X = CH₂, O, S, SO₂, CO, CMe₂
R = H, Me, OMe, OH

; optionally

substituted by one or more R^k selected from the group consisting of hydroxy, alkyl, alkoxy, alkoxyalkyl, thioalkyl, haloalkyl, cyano, amino, alkylamino, halogen, acylamino, sulfonamide and -COR;

R is hydroxy, alkoxy, alkyl or amino;

Z₁ is selected from the group consisting of CH₂, O, CH₂O, NH, CO, S, SO, CH(OH) and SO₂;

Z₂ is a 1-5 carbon linker optionally containing one or more heteroatom selected from the group consisting of O, S and N; or

Z₁ - Z₂ contains a moiety selected from the group consisting of carboxamide, sulfone, sulfonamide, alkenyl, alkynyl, and acyl; or

Z₁ - Z₂ contains a 5- or 6-membered aryl or heteroaryl ring optionally substituted with R^c, wherein the heteroaryl ring contains 1-3 heteroatoms selected from the group consisting of O, N and S;

R^c is selected from the group consisting of H, alkyl, haloalkyl, aryl, heteroaryl, halogen, alkoxyalkyl, aminoalkyl, hydroxy, alkoxy, carboxamide, and cyano;

wherein the carbon and nitrogen atoms of $Z_1 - Z_2$ are optionally substituted by a moiety selected from the group consisting of alkyl, alkoxy, thioalkyl, alkylsulfone, aryl, alkoxyalkyl, hydroxy, alkylamino, heteroaryl, alkenyl, alkynyl, carboxyalkyl, halogen, haloalky and acylamino;

B1 art.
cont.
X-Y contains a moiety selected from the group consisting of acyl, alkyl, sulfonyl, amino, ether, thioether, carboxamido, sulfonamido, aminosulfonyl and olefins; or

X is selected from the group consisting of $-CHR^c-$, $-NR^f-$, $-O-$, $-S-$, $-SO_2-$, and $-CO-$;

R^c is selected from the group consisting of H, lower alkyl, alkoxy, cycloalkyl, alkoxyalkyl, hydroxy, alkynyl, alkenyl, haloalkyl, thioalkyl and aryl; wherein when R^c is hydroxy, the hydroxy group can optionally form a lactone with the carboxylic acid function of the chain;

R^f is selected from the group consisting of H, alkyl, aryl, benzyl and haloalkyl;

Y is selected from the group consisting of $(CH_2)_p$, $-CHR^g-$, $-NR^g-$, CO and SO_2 ;


R^g is selected from the group consisting of H, alkyl, haloalkyl, alkoxyalkyl, alkynyl, aryl, heteroaryl, aralkyl, hydroxy, alkoxy, and carboxyalkyl;

p is 0 or 1;

Y^3 and Y^4 are independently selected from the group consisting of H, alkyl, haloalkyl, halogen, aryl, aralkyl, heteroaralkyl, heteroaryl, alkenes, hydroxyalkyl, and alkyne; wherein the alkyl chain is straight or branched and optionally contains one or more moieties selected from the group consisting of N, O, and S, sulfone,

sulfoamide, nitrile, carboxamide, carboalkoxy or carboxyl; wherein aryl and heteroaryl rings are monocyclic or bicyclic optionally containing 1-5 heteroatoms and wherein said ring may be saturated or unsaturated, and such rings may optionally be substituted by one or more substituent R^c ;

with the proviso that when Y^3 or Y^4 is H, Y^5 may be C or N, otherwise Y^5 is C; and

 R^b is $X_2 - R^h$ wherein X_2 is selected from the group consisting of O, S and NR^j wherein R^h and R^j are independently selected from the group consisting of H, alkyl, aryl, aralkyl, acyl and alkoxyalkyl.

Claim 72 (Original) A compound according to claim 71, wherein

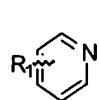
X is of $-CHR^e-$, wherein R^e is H;

Y is $-(CH_2)_p$ wherein $p = 0$;

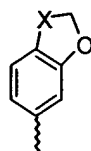
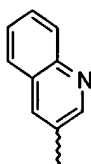
Y^5 is a carbon;

R^b is OH.

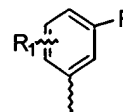
Claim 73 (Original) A compound according to Claim 72, wherein Y^3 or Y^4 is independently selected from the group consisting of:



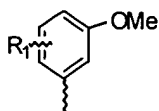
$R_1 = \text{H, alkyl, OMe, OH, halogen, amino, CN}$



$X = \text{CH}_2, \text{O}$



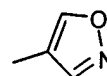
$R_1 = \text{H, alkyl, OMe, OH, halogen, amino, CN}$



$R_1 = \text{H, alkyl, OMe, OH, halogen}$

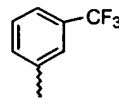
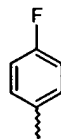
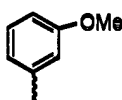
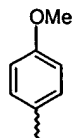
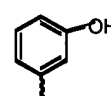
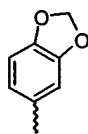
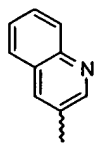
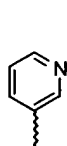


$X = \text{NH, NMe, O, S}$



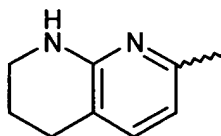
$\text{H, alkyl, CH}_2\text{B}_1\text{R}$ ($\text{B}_1 = \text{O, SO}_2, \text{S, CO; R} = \text{alkyl, aryl}$), CH_2OH ,
and $\text{---}\equiv\text{---R}$ ($\text{R} = \text{alkyl, aryl, alkoxyalkyl}$)

Claim 74 (Original) A compound according to claim 72, wherein Y^3 or Y^4 is independently selected from the group consisting of



$\text{H, Me, Ph, Et, Pr, i-Pr, and CH}_2\text{OCH}_2\text{Ph}$

Claim 75 (Original) A compound according to claim 74, wherein A^1 is



Claim 76 (Original) The method according to Claim 71 wherein the compound selected from the group consisting of

(2-{5-[3-(5,6,7,8-Tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-isoxazol-3-yl}-cyclopropyl)-acetic acid;
3-Phenyl-4-{5-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-isoxazol-3-yl}-butyric acid;
3-(2,3-Dihydro-benzofuran-6-yl)-4-{5-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-isoxazol-3-yl}-butyric acid;
3-(3-Fluoro-phenyl)-4-{5-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-isoxazol-3-yl}-butyric acid;
3-Pyridin-3-yl-4-{5-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-isoxazol-3-yl}-butyric acid;
3-Benzo[1,3]dioxol-5-yl-4-{5-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-isoxazol-3-yl}-butyric acid;
3-(2,3-Dihydro-benzofuran-6-yl)-4-{3-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-isoxazol-5-yl}-butyric acid;
3-(3-Fluoro-phenyl)-4-{3-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-isoxazol-5-yl}-butyric acid;
3-Pyridin-3-yl-4-{3-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-isoxazol-5-yl}-butyric acid;
3-Benzo[1,3]dioxol-5-yl-4-{3-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-isoxazol-5-yl}-butyric acid;
(2-{3-[3-(5,6,7,8-Tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-isoxazol-5-yl}-cyclopropyl)-acetic acid;
(2-{4-[3-(5,6,7,8-Tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-thiazol-2-yl}-cyclopropyl)-acetic acid;
3-Phenyl-4-{4-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-thiazol-2-yl}-butyric acid;
3-(2,3-Dihydro-benzofuran-6-yl)-4-{4-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-thiazol-2-yl}-butyric acid;
3-(3-Fluoro-phenyl)-4-{4-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-thiazol-2-yl}-butyric acid;

3-Pyridin-3-yl-4-{4-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-thiazol-2-yl}-butyric acid;

3-Benzo[1,3]dioxol-5-yl-4-{4-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-thiazol-2-yl}-butyric acid;

3-Phenyl-4-{3-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-isoxazol-5-yl}-butyric acid;

3-(2,3-Dihydro-benzofuran-6-yl)-4-{3-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-isoxazol-5-yl}-butyric acid;

3-(3-Fluoro-phenyl)-4-{3-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-isoxazol-5-yl}-butyric acid;

3-Pyridin-3-yl-4-{3-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-isoxazol-5-yl}-butyric acid;

3-Benzo[1,3]dioxol-5-yl-4-{3-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-isoxazol-5-yl}-butyric acid

(2-{5-[3-(5,6,7,8-Tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-isoxazol-3-yl}-cyclopropyl)-acetic acid;

3-Phenyl-4-{5-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-isoxazol-3-yl}-butyric acid;

3-(2,3-Dihydro-benzofuran-6-yl)-4-{5-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-isoxazol-3-yl}-butyric acid;

3-(3-Fluoro-phenyl)-4-{5-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-isoxazol-3-yl}-butyric acid;

3-Pyridin-3-yl-4-{5-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-isoxazol-3-yl}-butyric acid;

3-Benzo[1,3]dioxol-5-yl-4-{5-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-isoxazol-3-yl}-butyric acid;

3-(2,3-Dihydro-benzofuran-6-yl)-4-{3-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-isoxazol-5-yl}-butyric acid;

3-(3-Fluoro-phenyl)-4-{3-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-isoxazol-5-yl}-butyric acid;

3-Pyridin-3-yl-4-{3-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-isoxazol-5-yl}-butyric acid;

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3-Benzo[1,3]dioxol-5-yl-4-{3-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-isoxazol-5-yl}-butyric acid;
(2-{3-[3-(5,6,7,8-Tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-isoxazol-5-yl}-cyclopropyl)-acetic acid;
(2-{4-[3-(5,6,7,8-Tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-thiazol-2-yl}-cyclopropyl)-acetic acid;
3-Phenyl-4-{4-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-thiazol-2-yl}-butyric acid;
3-(2,3-Dihydro-benzofuran-6-yl)-4-{4-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-thiazol-2-yl}-butyric acid;
3-(3-Fluoro-phenyl)-4-{4-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-thiazol-2-yl}-butyric acid;
3-Pyridin-3-yl-4-{4-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-thiazol-2-yl}-butyric acid;
3-Benzo[1,3]dioxol-5-yl-4-{4-[3-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-propyl]-thiazol-2-yl}-butyric acid;
Phenyl-4-{3-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-isoxazol-5-yl}-butyric acid;
3-(2,3-Dihydro-benzofuran-6-yl)-4-{3-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-isoxazol-5-yl}-butyric acid;
3-(3-Fluoro-phenyl)-4-{3-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-isoxazol-5-yl}-butyric acid;
3-Pyridin-3-yl-4-{3-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-isoxazol-5-yl}-butyric acid; and
3-Benzo[1,3]dioxol-5-yl-4-{3-[2-(5,6,7,8-tetrahydro-[1,8]naphthyridin-2-yl)-ethoxy]-isoxazol-5-yl}-butyric acid.

Claim 77 (Original) A pharmaceutical composition comprising a therapeutically effective amount of a compound of claim 71 and a pharmaceutically acceptable carrier.

Claim 78 (Original) A pharmaceutical composition comprising a therapeutically effective amount of a compound of Claim 76 and a pharmaceutically acceptable carrier.

Claim 79 (Original) A pharmaceutical composition comprising a therapeutically effective amount of at least one compound of Claim 71 and a pharmaceutically acceptable carrier/or additive and optionally a cytotoxic agent.

Claim 80 (Original) A pharmaceutical composition comprising a therapeutically effective amount of at least one compound of Claim 76 and a pharmaceutically acceptable carrier/or additive and optionally a cytotoxic agent.



Claim 81 (Original) A method for treating conditions mediated by the $\alpha_v\beta_3$ integrin in a mammal in need of such treatment comprising administering an effective $\alpha_v\beta_3$ inhibiting amount of a compound of Claim 71.

Claim 82 (Original) A method for treating conditions mediated by the $\alpha_v\beta_3$ integrin in a mammal in need of such treatment comprising administering an effective $\alpha_v\beta_3$ inhibiting amount of a compound of Claims 76.

Claim 83 (Original) The method according to Claim 81 wherein the condition treated is tumor metastasis, solid tumor growth, angiogenesis, osteoporosis, humoral hypercalcemia of malignancy, smooth muscle cell migration, restenosis, atherosclerosis, macular degeneration, retinopathy, and arthritis.

Claim 84 (Original) The method according to Claim 82 wherein the condition treated is tumor metastasis, solid tumor growth, angiogenesis, osteoporosis, humoral hypercalcemia of malignancy, smooth muscle cell migration, restenosis, atherosclerosis, macular degeneration, retinopathy, and arthritis.
